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10/510,680 Youg Chu, 9-22-2006

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                 KOREAPAT updates resume
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NEWS 7 MAY 30
                 IPC 8 Rolled-up Core codes added to CA/CAplus and
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         MAY 30
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                 The first reclassification of IPC codes now complete in
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         JUN 02
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                 and display fields
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         JUl 14
NEWS 14 JUl 19
                 Coverage of Research Disclosure reinstated in DWPI
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                 INSPEC enhanced with 1898-1968 archive
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NEWS 16
        AUG 28
NEWS 17
         AUG 30
                 CA(SM)/CAplus(SM) Austrian patent law changes
NEWS 18
         SEP 11
                 CA/CAplus enhanced with more pre-1907 records
              JUNE 30 CURRENT WINDOWS VERSION IS V8.01b, CURRENT
NEWS EXPRESS
              MACINTOSH VERSION IS V6.0c(ENG) AND V6.0Jc(JP),
              AND CURRENT DISCOVER FILE IS DATED 26 JUNE 2006.
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=> file req SINCE FILE TOTAL COST IN U.S. DOLLARS ENTRY SESSION FULL ESTIMATED COST 0.21 0.21 CM 2

CRN 71-47-6 CMF C H O2

O== CH- O-

=> file reg

COST IN U.S. DOLLARS SINCE FILE TOTAL ENTRY SESSION FULL ESTIMATED COST 6.03 521.86

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L16 STRUCTURE UPLOADED

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Structure attributes must be viewed using STN Express query preparation.

=> s 116

SAMPLE SEARCH INITIATED 15:40:18 FILE 'REGISTRY'
SAMPLE SCREEN SEARCH COMPLETED - 1005 TO ITERATE

100.0% PROCESSED 1005 ITERATIONS

3 ANSWERS

55 ANSWERS

SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE \*\*COMPLETE\*\*

BATCH \*\*COMPLETE\*\*

PROJECTED ITERATIONS: 18199 TO 22001

PROJECTED ANSWERS:

3 TO 163

L17 3 SEA SSS SAM L16

=> s l16 full

FULL SEARCH INITIATED 15:40:26 FILE 'REGISTRY'
FULL SCREEN SEARCH COMPLETED - 19091 TO ITERATE

100.0% PROCESSED 19091 ITERATIONS

SEARCH TIME: 00.00.01

L18 55 SEA SSS FUL L16

=> file caplus

COST IN U.S. DOLLARS
SINCE FILE TOTAL
ENTRY SESSION
FULL ESTIMATED COST
167.38
689.24

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS) SINCE FILE TOTAL ENTRY SESSION

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$$R^2$$
 $R^1$ 
 $R^3$ 
 $CH_2)_n$ 
 $R^4$ 
 $R^5$ 
 $R^6$ 
 $Z-W-R$ 
 $I$ 

Title muscarinic receptor antagonists I (X = O, NH, etc.; R1 = OH, etc.; R2 = H, halo, alkyl; R3 = H, OH, etc.; R4, R5, R6 = H, alkyl; ; Z = CH2, SO2, carbonyl; W = alkylene, etc.; R = alkyl, aryl, etc.), useful for the treatment of various diseases of the respiratory, urinary and gastrointestinal systems mediated through muscarinic receptors, are prepd. The affinity of these compds. for M2 and M3 muscarinic receptor subtype was tested. For example, (3S)-1-benzylpyrrolidin-3-yl cyclopentyl(hydroxy)phenylacetate was prepd. and had pKi = 6.13/7.17 for the M2 and M3 receptor subtype resp.

TT 719278-65-6P 719278-66-7P 719278-72-5P
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(prepn. of 1-substituted-3-pyrrolidine derivs. as muscarinic receptor antagonists)

RN 719278-65-6 CAPLUS

CN Benzeneacetic acid, .alpha.-cyclopentyl-.alpha.-hydroxy-,
(3S)-1-[[4-(trifluoromethyl)phenyl]sulfonyl]-3-pyrrolidinyl ester (9CI)
(CA INDEX NAME)

Absolute stereochemistry.

RN 719278-66-7 CAPLUS

Absolute stereochemistry.

RN 719278-72-5 CAPLUS

### Absolute stereochemistry.

REFERENCE COUNT: 16 THERE ARE 16 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L19 ANSWER 2 OF 2 CAPLUS COPYRIGHT 2006 ACS on STN

Current application

ACCESSION NUMBER: DOCUMENT NUMBER:

2003:837081 CAPLUS 139:337885

TITLE:

Preparation of acyloxypyrrolidinium salts as M3

muscarinic antagonists

INVENTOR (S):

Prat Quinones, Maria; Fernandez Forner, Maria Dolors

Almirall Prodesfarma S.A., Spain

SOURCE:

PCT Int. Appl., 72 pp. CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT ASSIGNEE(S):

							DATE		APPLICATION NO.										
									WO 2003-EP3786										
WO	2003	0870	94		A3 20040318														
	W:	ΑE,	AG,	AL,	AM,	AT,	AU,	ΑZ,	BA,	BB	, BG,	BR,	BY,	BZ,	CA,	CH,	CN,		
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		FI,	FR,	GB,	GR,	HU,	ΙE,	IT,	LU,	MC.	, NL,	PT,	RO,	SE,	SI,	SK,	TR,		
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ES	2206021				A1 20040501				ES 2002-889						20020416				
ES	2206			B1 20050801															
CA	2482	536			AA 20031023			CA 2003-2482536						20030411					
AU	U 2003233967					A1 20031027				AU 2003-233967						20030411			
EP	1497			A2 20050119				EP 2003-727294						20030411					
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		ΙE,	SI,	LT,	LV,	FI,	RO,	MK,	CY,	AL	, TR,	ВG,	CZ,	EE,	HU,	SK			
BR	2003	0091	67		A 20050125				BR 2003-9167						20030411				
	1662						2005	0831	CN 2003-813892						20030411				
ZA	ZA 2004008335						A 20051102			ZA 2004-8335						20041014			
NO	NO 2004004826						A 20050114			NO 2004-4826						20041105			
US	US 2005282875						A1 20051222			US 2005-510680						20050720			
PRIORITY	PRIORITY APPLN. INFO.:									ES 2	2002-	889		7	A 2	0020	416		
							1	WO :	2003-1	EP37	86	I	<i>N</i> 2	0030	411				

OTHER SOURCE(S):

MARPAT 139:337885

GI

$$R-(CH_2)_{n}-A-(CH_2)_{m}-N$$
 $X^{-}$ 
I

$$\begin{array}{c|c} & & & \\ & & & \\$$

AB Pyrrolidinium derivs. I [R = (un)substituted Ph, naphthalenyl, 5,6,7,8-tetrahydronaphthalenyl, benzo[1,3]dioxolyl, biphenyl, heteroarom.; R1 = alkyl; R2 = CR3R4R5, Q; R3 = 2-furyl, 3-furyl, 2-thienyl, 3-thienyl; R4 = 2-furyl, 3-furyl, 2-thienyl, 3-thienyl, cycloalkyl; R5 = H, OH, Me, CH2OH; Q1 = CH2, CH2CH2, O, OCH2, S, SCH2, CH:CH; A = (un)substituted CH:CH, CH2, CO, O, S, S(O), SO2, NH; m = 0-8; n = 0-4] were prepd. for use in therapy as antagonists of M3 muscarinic receptors (no data). Thus, (3R)-3-pyrrolidinol was treated with 2-(3-bromopropyl)thiophene to give (3R)-1-(3-thien-2-ylpropyl)pyrrolidinol which was treated with Me 2-hydroxy-2,2-dithen-2-ylacetate and quaternized to give the pyrrolidinium salt II.

II

IT 616865-64-6P 616865-65-7P
RL: PUR (Purification or recovery); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(prepn. of acyloxypyrrolidinium salts as M3 muscarinic antagonists) 616865-64-6 CAPLUS

CN Pyrrolidinium, 3-[(hydroxydi-2-thienylacetyl)oxy]-1-methyl-1-(3-phenoxypropyl)-, bromide, (1S,3R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN

RN 616865-65-7 CAPLUS
CN Pyrrolidinium, 3-[(hydroxydi-2-thienylacetyl)oxy]-1-methyl-1-(3-phenoxypropyl)-, bromide, (1R,3R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

Br -

Absolute stereochemistry.

CM 1

CRN 616866-05-8 CMF C22 H23 N O4 S2

CM 2

CRN 144-62-7 CMF C2 H2 O4

RN 616866-07-0 CAPLUS
CN 2-Thiopheneacetic acid, .alpha.-hydroxy-.alpha.-2-thienyl-,
(3R)-1-(3-phenoxypropyl)-3-pyrrolidinyl ester (9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 616866-08-1 CAPLUS

CN 2-Thiopheneacetic acid, .alpha.-hydroxy-.alpha.-2-thienyl-, (3R)-1-(3-phenoxypropyl)-3-pyrrolidinyl ester, ethanedioate (1:1) (salt) (9CI) (CA INDEX NAME)

CM 1

CRN 616866-07-0 CMF C23 H25 N O4 S2

Absolute stereochemistry.

CM 2

CRN 144-62-7 CMF C2 H2 O4

IT 616865-58-8P 616865-59-9P 616865-60-2P 616865-62-4P 616865-63-5P 616865-76-0P 616865-77-1P 616865-78-2P 616865-86-2P

Absolute stereochemistry.

#### ● Br~

RN 616865-59-9 CAPLUS
CN Pyrrolidinium, 3-[(hydroxydi-2-thienylacetyl)oxy]-1-methyl-1-(2-phenoxyethyl)-, bromide, (1R,3R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

#### ● Br -

RN 616865-60-2 CAPLUS
CN Pyrrolidinium, 3-[(hydroxydi-2-thienylacetyl)oxy]-1-methyl-1-(2-phenoxyethyl)-, bromide, (1S,3R)- (9CI) (CA INDEX NAME)

• Br-

RN 616865-62-4 CAPLUS
CN Pyrrolidinium, 3-[(hydroxydi-2-thienylacetyl)oxy]-1-methyl-1-(3-phenoxypropyl)-, bromide (9CI) (CA INDEX NAME)

• Br-

RN 616865-63-5 CAPLUS
CN Pyrrolidinium, 3-[(hydroxydi-2-thienylacetyl)oxy]-1-methyl-1-(3-phenoxypropyl)-, bromide, (3R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

• Br-

RN 616865-76-0 CAPLUS
CN Pyrrolidinium, 3-[(cyclohexyl-2-furanylhydroxyacetyl)oxy]-1-methyl-1-[2-(phenylmethoxy)ethyl]-, bromide, (3R)- (9CI) (CA INDEX NAME)

# ● Br-

RN 616865-77-1 CAPLUS

CN Pyrrolidinium, 3-[[(2R)-cyclopentylhydroxyphenylacetyl]oxy]-1-methyl-1-[2-(phenylmethoxy)ethyl]-, bromide, (1R,3R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

# • Br-

RN 616865-78-2 CAPLUS

CN Pyrrolidinium, 3-[[(2R)-cyclopentylhydroxyphenylacetyl]oxy]-1-methyl-1-[2-(phenylmethoxy)ethyl]-, bromide, (1S,3R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

## • Br-

RN 616865-86-2 CAPLUS

CN Pyrrolidinium, 1-[3-(2-benzothiazolyloxy)propyl]-3-[[(2R)-cyclohexyl-2furanylhydroxyacetyl]oxy]-1-methyl-, chloride, (3R)- (9CI) (CA INDEX NAME)

$$\begin{array}{c} \text{Me} \\ \text{N} \\ \text{N} \end{array}$$

● c1 -

RN 616865-87-3 CAPLUS

CN Pyrrolidinium, 1-[3-(2-benzothiazolyloxy)propyl]-3-[[(2S)-cyclohexyl-2furanylhydroxyacetyl]oxy]-1-methyl-, chloride, (3R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

$$\begin{array}{c|c} & & & \\ &$$

• c1-

RN 616865-88-4 CAPLUS

CN Pyrrolidinium, 1-[3-(2-benzothiazolyloxy)propyl]-3-[(cyclohexyl-2-furanylhydroxyacetyl)oxy]-1-methyl-, chloride, (3R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

$$\begin{array}{c|c}
 & Me \\
 & N \\$$

● Cl -

RN 616865-89-5 CAPLUS

CN Pyrrolidinium, 3-[[(2R)-cyclohexyl-2-furanylhydroxyacetyl]oxy]-1-ethyl-1-

[3-(phenylthio)propyl]-, bromide, (3R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

● Br -

RN 616865-90-8 CAPLUS

CN Pyrrolidinium, 3-[[(2S)-cyclohexyl-2-furanylhydroxyacetyl]oxy]-1-ethyl-1-[3-(phenylthio)propyl]-, bromide, (3R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

● Br -

RN 616865-91-9 CAPLUS

CN Pyrrolidinium, 3-[[(2R)-cyclopentylhydroxyphenylacetyl]oxy]-1-methyl-1-[3-(phenylthio)propyl]-, bromide, (1R,3R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

● Br -

RN 616865-92-0 CAPLUS

CN Pyrrolidinium, 3-[[(2R)-cyclopentylhydroxyphenylacetyl]oxy]-1-methyl-1-[3-(phenylthio)propyl]-, bromide, (1S,3R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

• Br-

RN 616865-93-1 CAPLUS

CN Pyrrolidinium, 3-[[(2R)-cyclopentylhydroxyphenylacetyl]oxy]-1-methyl-1-[3[(5,6,7,8-tetrahydro-2-naphthalenyl)oxy]propyl]-, bromide, (1S,3R)- (9CI)
(CA INDEX NAME)

Absolute stereochemistry.

• Br-

RN 616865-94-2 CAPLUS

CN Pyrrolidinium, 3-[[(2R)-cyclopentylhydroxyphenylacetyl]oxy]-1-methyl-1-[3-[(5,6,7,8-tetrahydro-2-naphthalenyl)oxy]propyl]-, bromide, (1R,3R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

• Br-

RN 616865-95-3 CAPLUS

CN Pyrrolidinium, 3-[(2R)-cyclopentylhydroxyphenylacetyl]oxy]-1-[3-(4-methoxyphenoxy)propyl]-1-methyl-, bromide, (3R)- (9CI) (CA INDEX NAME)

• Br

RN 616865-96-4 CAPLUS

CN Pyrrolidinium, 3-[[(2R)-cyclopentylhydroxyphenylacetyl]oxy]-1-[3-(4-methoxyphenoxy)propyl]-1-methyl-, bromide, (3S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

● Br-

RN 616866-00-3 CAPLUS

CN Pyrrolidinium, 3-[(hydroxydi-2-thienylacetyl)oxy]-1-[3-(3-hydroxyphenoxy)propyl]-1-methyl-, (3S)-, formate (9CI) (CA INDEX NAME)

CM 1

CRN 616865-99-7 CMF C24 H28 N O5

CMF C24 H28 N O5 S2

Absolute stereochemistry.

CM 2

CRN 71-47-6 CMF C H O2 RN 616866-02-5 CAPLUS
CN Pyrrolidinium, 1-[3-(3-cyanophenoxy)propyl]-3-[(cyclohexyl-2-furanylhydroxyacetyl)oxy]-1-methyl-, (3R)-, formate (9CI) (CA INDEX NAME)

CM 1

CRN 616866-01-4
CMF C27 H35 N2 O5

Absolute stereochemistry.

CM 2

CRN 71-47-6 CMF C H O2

O== CH-O-

RN 616866-04-7 CAPLUS
CN Pyrrolidinium, 3-[(cyclohexyl-2-furanylhydroxyacetyl)oxy]-1-methyl-1-[3-(1-naphthalenyloxy)propyl]-, (3R)-, formate (9CI) (CA INDEX NAME)

CM 1

CRN 616866-03-6 CMF C30 H38 N O5

CRN 71-47-6 CMF C H O2

O CH-O

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L19 2 L18

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L19 ANSWER 1 OF 2 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2004:546475 CAPLUS

DOCUMENT NUMBER:

141:106362

TITLE:

Preparation of 1-substituted-3-pyrrolidine derivatives

as muscarinic receptor antagonists

INVENTOR (S):

Mehta, Anita; Gupta, Jang Bahadur; Sarma, Pakala

Kumara Savithru

PATENT ASSIGNEE(S):

Ranbaxy Laboratories Limited, India

SOURCE:

PCT Int. Appl., 47 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

LANGUAGE:

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

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IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, SK US 2006194862 PRIORITY APPLN. INFO.:

OTHER SOURCE(S):

A1 20060831 US 2006-540245 WO 2002-IB5590

CASREACT 141:106362; MARPAT 141:106362

GI